## Mechanisms of Atiulcerogenic Activity of Costus speciosus Rhizome Extract in Ethanol-Induced Gastric Mucosal Injury in Rats

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Abstract : Costus speciosus is an important Malaysian medicinal plant commonly used traditionally in the treatment of many aliments. The present investigation is designed to elucidate preventive effects of ethanolic extracts of C. speciosus rhizome against absolute ethanol-induced gastric mucosal injury in Sprague-Dawley rats. Five groups of rats were orally pre-treated with vehicle, carboxymethylcellulose (CMC) as normal control group (Group 1), ethanol as ulcer control group (Group 2), omeprazole 20 mg/kg (reference group) (Group 3), and 250 and 500 mg/kg of C. speciosus extract (experimental groups) (Group 4 and 5), respectively. An hour later, CMC was given orally to Group 1 rats and absolute ethanol was given orally to Group 2-5 rats to generate gastric mucosal injury. After an additional hour, the rats were sacrificed. Grossly, ulcer control group exhibited severe of gastric mucosal hemorrhagic injury and increased in ulcer area, whereas groups pre-treated with omeprazole or plant's rhizomes exhibited the significant reduction of gastric mucosal injury. Significant increase in the pH and mucous of gastric content was observed in rats re-treated with C. speciosus rhizome. Histology, ulcer control rats, demonstrated remarkable disruption of gastric mucosa, increased in edema and inflammatory cells infiltration of submucosal layer compared to rats pre-treated with rhizomes extract. Periodic acid Schiff staining for glycoprotein, rats pre-fed with C. speciosus C. displayed remarkably intense uptake of magenta color by glandular gastric mucosa compared with ulcer control rats. Immunostaining of gastric epithelium, rats pre-treatment with rhizome extract provide evidence of up-regulation of HSP70 and down-regulation of Bax proteins compared to ulcer control animals. Gastric tissue homogenate, C. speciosus significantly increased the activity of superoxide dismutase (SOD), and catalase (CAT), increased the level of non-protein sulfhydryl (NP-SH) and decreased the level of lipid peroxidation after ethanol administration. Acute toxicity test did not show any signs of toxicity. The mechanisms implicated the gasrtoprotective property of C. speciosus depend upon the antisecretory activity, increased in gastric mucus glycoprotein, up-regulation of HSP70 protein and down-regulation of Bax proteins, reduction in the lipid peroxidation and increase in the level of NP-SH and antioxidant enzymes activity in gastic homogenate. Keywords : antioxidant, Costus speciosus, gastric ulcer, histology, omeprazole

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1

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